

Please Direct All Correspondence to Customer Number **30,313**

**ISSUE FEE TRANSMITTAL LETTER**

Applicant : Audrey GODDARD et al.  
App. No : 10/063,602  
Filed : May 3, 2002  
For : SECRETED AND  
TRANSMEMBRANE  
POLYPEPTIDES AND NUCLEIC  
ACIDS ENCODING THE SAME  
Art Unit : 1647  
Class/Sub-Class : 435-069100  
Examiner : WEGERT, Sandra L.

**CERTIFICATE OF MAILING**

I hereby certify that this correspondence and all marked attachments are being deposited with the United States Postal Service as first-class mail in an envelope addressed to: Mail Stop Issue Fee, Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450, on

February 26, 2007

(Date)

  
Kerry Taylor, Reg. No. 43,947

**MAIL STOP ISSUE FEE**

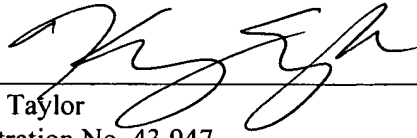
Commissioner for Patents  
P.O. Box 1450  
Alexandria, VA 22313-1450

Dear Sir:

Enclosed for filing is the Issue Fee for the above-identified application:

- (X) Form PTOL-85.
- (X) Letter Regarding Patent Term Adjustment Calculation in 1 page.
- (X) Comments on Statement of Reasons for Allowance in 3 pages.
- (X) A check in the amount of \$1,730 is enclosed for the following fees:
  - (X) \$1,400 Issue Fee
  - (X) \$300 Publication Fee
  - (X) \$30 Advance Order of 10 Copies
- (X) Return prepaid postcard.

The Commissioner is hereby authorized to charge any additional fees which may be required, or credit any overpayment, to Account No. 11-1410.

  
Kerry Taylor  
Registration No. 43,947  
Attorney of Record  
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## IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant : Goddard, *et al.*  
Appl. No. : 10/063,602  
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For : SECRETED AND  
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February 26, 2007

(Date)

  
Kerry Taylor, Reg. No. 43947

COMMENTS ON STATEMENT OF REASONS FOR ALLOWANCE

Mail Stop Issue Fee  
Commissioner for Patents  
P.O. Box 1450  
Alexandria, VA 22313-1450

Dear Sir:

Applicants received a Notice of Allowability and Examiner's Statement of Reasons for Allowance mailed from the USPTO on November 27, 2006. Applicants submit the following Comments on the Examiner's Statement.

The Examiner's Statement includes the following:

Applicant relies on more than 140 references (see IDS filed 7/7/06 and 9/14/06), where expression levels of mRNA, measured by quantitative PCR, were found to have a good correlation to the expressed protein levels. .... While the PTO found several references in which the protein expression levels did not correlate with mRNA levels measured by quantitative PCR, the majority of the references which were found, including those cited by Applicant, demonstrated a correlation between mRNA levels measured by quantitative PCR and protein expression levels.

Applicants assert that the expression levels of protein correlate to mRNA (cDNA) levels when the cDNA is measured by quantitative PCR (i.e. rtPCR). Applicant has provided more than 140 references in support of this position. *Notice of Allowability* at 4-5 (emphasis added, citations omitted).

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**Filed** : **May 3, 2002**

Applicants have submitted as exhibits numerous references in support of their assertion that, generally speaking, differential mRNA expression levels (e.g., tumor tissue vs. normal tissue) result in similar differential expression levels of the encoded protein. In addition, Applicants have submitted several Information Disclosure Statements listing numerous additional references, including those submitted as exhibits. Applicants have not represented to the PTO that all of the references submitted to the PTO in an IDS support Applicants' assertions.

Applicants have also argued that statements by the authors of Hu *et al.* (J. Proteome Res. 2003; 2(4):405-12) regarding the significance of differential mRNA expression detected by mRNA microarrays are not relevant to the data relied on by Applicants. Applicants' data in Example 18 of the instant application are based on RT-PCR, which is recognized by those of skill in the art as more sensitive and reliable than mRNA microarrays, and therefore the opinions of Hu regarding the significance of differential mRNA data from microarrays is not relevant to the instant application. Applicants' have not argued that microarray data are not reliable, or that all of the supporting references relied on by the Applicants used quantitative RT-PCR to measure mRNA levels.

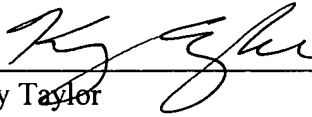
Finally, Applicants note for the record that they do not necessarily agree with the PTO's characterization of the references cited in the Examiner's statement of reasons for allowance. Applicants do agree with the PTO's conclusion that "[b]ased on the totality of evidence of record, one of skill in the art would find it more likely than not that an increase in message as measured by rtPCR would be predictive of an increase in protein expression levels, absent evidence to the contrary." *Notice of Allowability* at 5.

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Filed : May 3, 2002

Respectfully submitted,

KNOBBE, MARTENS, OLSON & BEAR, LLP

Dated: 26-Feb-07

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